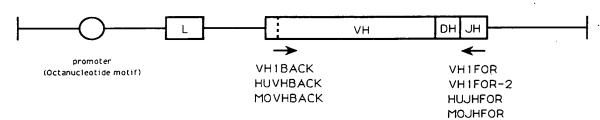


Inventor: Gregory P. WINTER,

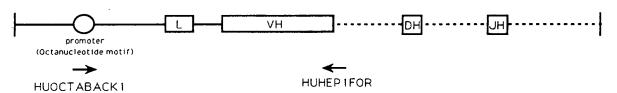
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6545142

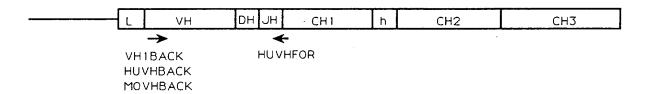




Unrearranged heavy chain variable gene (DNA)



Rearranged heavy chain variable gene (mRNA)



Rearranged light chain variable gene (DNA).

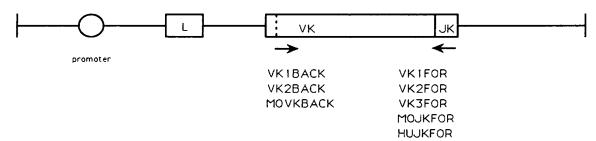
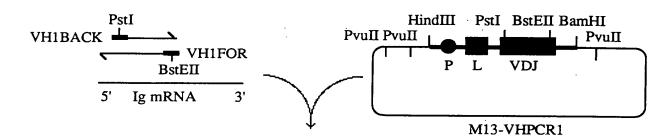


FIG. 1



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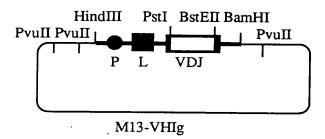


FIG. 2

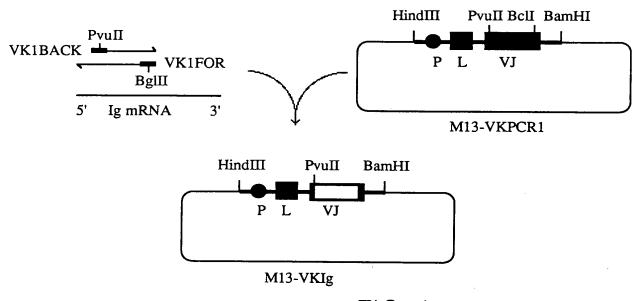


FIG. 4



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M13 VHPCR1.

HinD III(1)					·
AAGCTTATGAATAT	GCAAATCCTCTG 20	AATCTACATO	GTAAATATA 40	GGTTTGTCTA' 50	TACCA 60
CAAACAGAAAAACA' 70	TGAGATCACAGT 80	TCTCTCTAC	AGTTACTGAG 100	CACACAGGAC	CTCAC 120
M G W S (F L V A		aggtaagggg	CTCAC
130	140	150	160	170	180
AGTAGCAGGCTTGA 190	GGTCTGGACATA 200	TATATGGGT0 210	GACAATGACA 220	rccactttgc 230	CTTTC 240
G V			ESG		
TCTCCACAGGTGTC	CACTCCCAGGTC 260	CAA <u>CTGCAG</u> 270	GAGAGCGGTC 280	CAGGTCTTGT 290	GAGAC 300
15 S Q T L	SLTC	25 T V S	G S T		CDR1 W M
CTAGCCAGACCCTG	AGCCTGACCTGC 320	ACCGTGTCT(330	GCAGCACCT 340	350	360
35 <i>H</i> W V R	40 Q P P G	45 R G L	E W I	50 G R I D	DR2
TGCACTGGGTGAGA 370	380	390	400	410 70	420
55 S G G T ATAGTGGTGGTACT 430	60 K Y N E AAGTACAATGAG 440	65 K F K AAGTTCAAG 450	S R V AGCAGAGTGA 460	T M L V	D T AGACA 480
75 S K N Q CCAGCAAGAACCAG	80 F S L R	85 L S S	V T A	90 A D T A	
490	500 CDR3	510	520	530	540
95 Y C A R ATTATTGTGCAAGA 550	100 Y D Y Y			110 Y W G Q ACTGGGGCCA 590	
BstEII 115	120	370			
T V T V CCAC <u>GGTCACCG</u> TC 610	SS	TCCTTACAA	CCTCTCTCTT 640	CTATTCAGCT 650	TAAAT 660
AGATTTTACTGCAT 670	TTGTTGGGGGGG	GAAATGTGTG 690	TATCTGAATT 700	TCAGGTCATG 710	AAGGA 720
CTAGGGACACCTTG 730	GGAGTCAGAAAG 740	GGTCATTGG 750	GAGCCCGGGC 760	TGATGCAGAC 770	AGACA 780
			BamHI 		
TCCTCAGCTCCCAG 790	ACTTCATGGCCA 800	GAGATTTAT. 810	A <u>G</u>	FIG.	3



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M13 VkPCR1 "HinD III AAGCTTATGAATATGCAAATCCTCTGAATCTACATGGTAAATATAGGTTTGTCTATACCA 58 68 48 CAAACAGAAAAACATGAGATCACAGTTCTCTCTACAGTTACTGAGCACACAGGACCTCAC 128 118 108 MGWSCIILFLVATAT CATGGGATGGAGCTGTATCATCCTCTTCTTGGTAGCAACAGCTACAGGTAAGGGGCTCAC 158 168 178 188 198 208 AGTAGCAGGCTTGAGGTCTGGACATATATATGGGTGACAATGACATCCACTTTGCCTTTC 238 248 258 268 228 218 Pvu II 10 G V H S D I Q L T Q S P S S L S A S 318 328 308 278 288 CDR1 25 20 V G D R V T I T C R A S G N I H N Y L A GCGTGGGTGACAGAGTGACCATCACCTGTAGAGCCAGCGGTAACATCCACAACTACCTGG 348 358 368 378 338 CDR2 45 50 40 W Y Q Q K P G K A P K L L I Y Y T T T L CTTGGTACCAGCAGAAGCCAGGTAAGGCTCCAAAGCTGCTGATCTACTACACCACCACCC 408 418 428 438 60 65 ADGVPSRFSGSGSGTDFTFT TGGCTGACGGTGTGCCAAGCAGATTCAGCGGTAGCGGTAGCGGTACCGACTTCACCTTCA 468 478 488 498 458 CDR3 90 85 80 ISSLQPEDIATYYC*QHFWST* CCATCAGCAGCCTCCAGCCAGAGGACATCGCCACCTACTACTGCCAGCACTTCTGGAGCA 528 538 548 558 Bcl I (requires dam host) 100 105 108 PRTFGQGTKVVIKR CCCCAAGGACGTTCGGCCAAGGGACCAAGGTGG<u>TGATCA</u>AACGTGAGTAGAATTTAAACT 578 588 598 608 BamHI

TTGCTTCCTCAGTTGGATCC

648

638



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Sequence of MBrl VH

Splice ↓G V H S AGGTGTCCACTCC 10 20 PstI Q V Q L Q E S G T E L A S P G A S V T L CAGGTCCAACTGCAGGAGTCAGGAACTGAGCTGGCGAGTCCTGGGGCATCAGTGACACTG VH1BACK SITE 30 <u>CDR1</u> 40 S C K A S G Y T F T <u>D H I I N</u> W V K K R TCCTGCAAGGCTTCTGGCTACACATTTACTGACCATATTATAAATTGGGTAAAAAAGAGG 52a 53 CDR2 PGQGLEWIGRI 70 60 CDR2 65 NOKFMGKATFSVDRSSNTVY AATCAAAAATTCATGGGCAAGGCCACATTCTCTGTAGACCGGTCCTCCAACACAGTGTAC 80 82A B C 83 90 MVLNSLTSEDPAVYYCGRGF ATGGTGTTGAACAGTCTGACATCTGAGGACCCTGCTGTCTATTACTGTGGAAGGGGCTTT CDR3 103 BstEII Splice D F D Y W G Q G T T V T V S S L GATTTTGACTACTGGGGCCAAGGGACCACGGTCACCGTCTCCTCAGGT..... VH1FOR SITE

Sequence of MBrl VK

Splice -1 ↓G V H S AGGTGTCCACTCC PvuII 10 D I Q L T Q S P P S L T V S V G E R V T GACATTCAGCTGACCCAGTCTCCACCATCCCTGACTGTGTCAGTAGGAGAGAGGGTCACT VK1BACK SITE <u> 27A</u> I S C K S N Q N L L W S G N R R Y C L G ATCAGTTGCAAATCCAATCAGAATCTTTTATGGAGTGGAAACCGAAGGTACTGTTTGGGC 40 50 CDR2 35 W H Q W K P G Q T P T P L I T W T S D R TGGCACCAGTGGAAACCAGGGCAAACTCCTACACCGTTGATCACCTGGACATCTGATAGG 60 FSGVPDRFIGSGSVTDFTLT TTCTCTGGAGTCCCTGATCGTTTCATAGGCAGTGGATCTGTGACAGATTTCACTCTGACC 80 90 CDR3 I S S V Q A E D V A V Y F C Q Q H L D L ATCAGCAGTGTGCAGGCTGAAGATGTGGCAGTTTATTTCTGTCAGCAACATTTGGACCTT 100 BglII/BclI Splice PYTFGGGTKL<u>EI</u>K↓ CCGTACACGTTCGGAGGGGGGGCCAAGCTGGAGATCAAACGTGAG VK1FOR SITE



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α -Lys 30

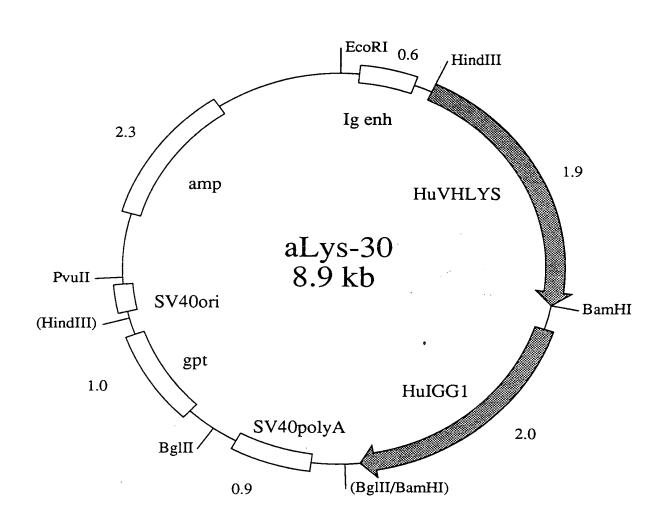
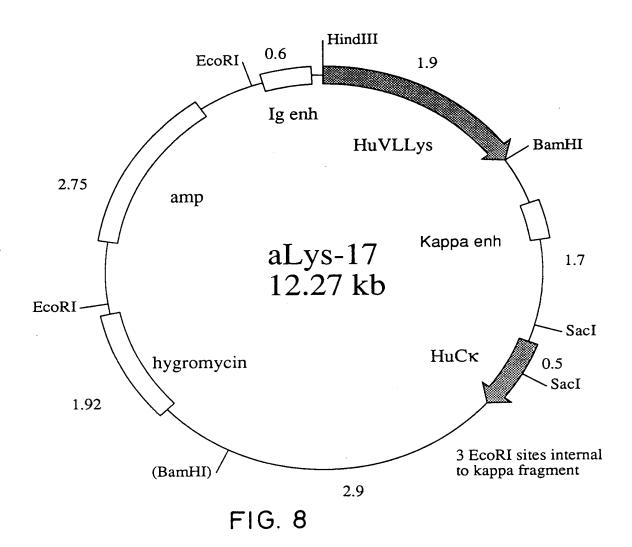


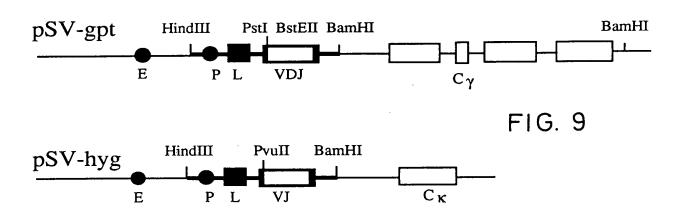
FIG. 7



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α -Lys 17







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	FR1	CDR_1	FR2	CDR 2
KABAT	Γ IA			
A07 A09 E03 G01	PGLVKPSQSLSLTCSVTGYSIT PGLVKPSQSLFLTCSITGFPIT PGLVKPSQSLSLTCSVTGYSIT PGLVKPSQSLSLTCSVTGYSIT	SGYYWN SGYYWI SGYYWN SGYYWN	WIRQFPGNKLEWMG WIRQSPGKPLEWMG WIRQFPGNKLEWMG WIRQFPGNKLEWMG	YISYDGSNNYNPSLKN YITHSGETFYNPSLQS YISYDGSNNYNPSLKN YISYDGSNNYNPSLKN
KABAT	T IB			
A06 25G07 B03 G03 H09 25C10 A12 A08 25G08 A03 C07 H04	PVLVAPSQSLSITCAVSDFSLT PGLVQPSQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PGLVQPSQSLSITCTVSGFSLT PVLVAPPQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PGLVAPSQSLSITCTVSGFSLT PVLVAPSQSLSITCTVSGFSLT PVLVAPSQSLSITCTVSGFSLT	NYGVL SYGVH SYGVH SYGVH SYAIS SYAIS SYGVH SYDVD SYGVH SYGVH SYGVH	WVRQPPGKGLEWLG WVRQSPGKGLEWLG WVRQPPGKGLEWLG WVRQPPGKGLEWLG WVRQPPGKGLEWLG WVRQPPGKGLEWLG WVRQPPGKGLEWLG WVRQPPGKGLEWLG WVRQSPGKGLEWLG WVRQSPGKGLEWLG WVRQSPGKGLEWLG WVRQSPGKGLEWLG	VIWAGGITNYNSALMS VIWSGGSTDYNAAFIS VIWGGGSTNYNSALMS VIWSGGSTDYNAAFIS VIWAGGSTNYNSALMS VIWTGGGTNYNSALKS VIWTGGGTNYNSALKS VIWGGGSTNYNSALKS VIWGGGSTNYNSALKS VIWSGGSTDYNAAFIS VIWAGGSTNYNSALMS VIWAGGSTNYNSALMS
KABAT	' IIA			
E04 H07	PELVRPGVSVKISCKGSGYTFT PELVRPGVSVKISCKGSGYTFT	DYAMH DYAMH	WVKQSHAKSLEWIG WVKQSHAKSLEWIG	VISTYYGDASYNQKFKD VISTYYGDASYNQKFKD
KABAT	IIB			
A02 B04 C05 C09 D06 D08 E07 G08 G10 25G09 F04 H02 H01 25C05 B01 B05 B11	AELVMPGASVKLSCKASGYTFT AELVKPGASVKMSCKASGYTFT AELVKPGASVKLSCKASGYTFT AELVKPGASVKLSCKASGYTFT ASLVKPGASVKLSCKASGYTFT PELVKPGASVKLSCKASGYTFT AELVRPGASVKLSCKASGYTFT AELVRPGASVKLSCKASGYTFT AELVKPGASVKVSCKASGYTFT AELVKPGASVKVSCKASGYTFT TELVKPGASVKLSCKASGYTFT TELVKPGASVKLSCKASGYTFT AELVKPGASVKLSCKASGYTFT AELVKPGASVKLSCKASGYTFT AELVKPGASVKLSCKASGYTFT AELVKPGASVKLSCKASGYTFT AELVRPGSSVKLSCKASGYTFT AELVRPGSSVKLSCKASGYTFT AELVRPGSSVKLSCKASGYTFT AELVRPGSSVKLSCKASGYTFT AELVRPGSSVKLSCKASGYTFT AELVRPGSSVKLSCKASGYTFT	SYWMH SYWIT SYWMH SYWIT SYWMH DYYIN SYWMH TYPIE SYWMH SYWMH SYWMH NYWMK SYWIT RHAMH SYWIT	WVKQRPGQGLEWIG	EIDPSDSYTNYNOKFKG DIYPGSGSTNYNEKFKS RIDPNSGGTKYNEKFKS EINPSNGGTNYDEKFKS DIYPGSGSTNYNEKFKS EINPSNGGTNYNEKFKS EINPSNGGTNYNEKFKS AIDPETGGTAYNOKFKG WIYPGSGNTKYNEKFKG RIHPSDSDTNYNOKFKG NFHPYNDDTKYNEKFKG NINPSNGGTNYNOKFKG NIDPSNGGTNYNOKFKD EIDPSDSYTNYN*KVQG QIFPASGSIYYNEMHKD DIYPGSGSTNYNEKFKS SFTMYSDATEYSENFKG DIYPGSGSTNYNEKFKS
KABAT	III A			
25G05 C10 B07	GGLVQAWGSLSLSCAASGFTFT GGLVQPGGSLSLSCAASGFTFT GGLVQPGGSLSLSCAASGFTFT	DYYMS DYYMN DYYMS	WVRQPPGKALEWLG WVRQPPGKALEWLA WVRQPPGKALEWLA	FIRNKANGYTTEYSASVKG LIRHKANGYTMEYSASVKG LIRNKANGYTTEYSASVKG
KABAT	ІІІ В			
G05 B12 D04 D05 F12 F06 D02 F09	GGLVKPGGSLKLSCAASGFTFS GGLVQPGESLKLSCESNEYEFP GGLVQPGGSLRLSCAASGFTFS GGLVQPGGSLRLSCAASGFTFS GGLVQPGESWKLSCVIQQ**** GGLVQPGGSLRLSCAASGFTFS GGLVQPGGSLRLSCAASGFTFS GGLVQPGESLKLSCESNEYVIP GDLVKPGGSLKLSCAASGFTFS	DYGMH SHDMS SYAMS SYAMS ***** SYAMS *HDMS SYGMS	WVRQAPEKGLEWVA WVR********** WVA*APGKGLEWVS WVA*APGKGLEWVS WVRQ*PEKRLELVA WVA*APGKGLEWVS WVRQTPGLEWVS	YISSGSSTIYYADTVKG AINSDGGSTYYPDTMER AISGSGGSTYYADSVKG AISGSGGSTYYADSVKG AINSDGGSTYYPDTMER AISGSGGSTYYADSAKG AINSDGGSTYYPDTMER TISSGGSYTYYPDSVKG
KABAT	ш с			
E06	GGLVQPGGSMKLSCAASGFTFS	DAWMD	WVRQSPEKGLEWVA	EIRNKANNHATYYAESVKG
KABAT	V A			
C04	AELVKPGASVKLSCKASGYTFT	EYTIH	WVKORSGOGLEWIG	WFYPGSGSIKYNEKEKD

C04 AELVKPGASVKLSCKASGYTFT EYTIH WVKQRSGQGLEWIG WFYPGSGSIKYNEKFKD





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CDR 3

RISITRDTSKNQFFLKLNSVTTEDTATYYCAR PISITRETSKNQFFLQLNSVTTEDTAMYYCAG RISITRDTSKNQFFLQLNSVTTEDTATYYCAR

RISITRDTSKNQFFLKLNSVTTEDTATYYCAR

FR 3

EGNWDGFAY DRDKLGPWFAY DSSGSMDY VSSGYESMDY

RLSISKDTSKSQVFLKMNSLQTDDTAVYYCAK
RLSISKDNSKSQVFFKMNSLQADDTAIYYCAR
RLSISKDNSKSQVFFKMNSLQADDTAIYYCAR
RLSISKDNSKSQVFFKMNSLQADDTAIYYCAR
RLSISKDNSKSQVFLKMNSLQTDDTAMYYCAI
RLSISKDNSKSQVFLKMNSLQTDDTARYYCAR
RLSISKDNSKSQVFLKMNSLQTDDTARYYCAR
RLSISKDNSKSQVFLKMNSLQTDDTAMYYCAR
RLSISKDNSKSQVFLKMNSLQTDDTAMYYCAR
RLSISKDNSKSQVFFKMNSLQADDTAIYYCAR
RLSISKDNSKSQVFLKMNSLQTDDTAMYYCAR
RLSISKDNSKSQVFLKMNSLQTDDTAMYYCAK
RLSISKDNSKSQVFLKMNSLQTDDTAMYYCAK

HGDSSGYFDY NDGYY LGRGYAMDY KRDYDYDRGYYYAMDY YYDGSFFAY EGYYYFAY IYYDGSSDYYAMDY

13 nt. Ps.gene/Unproductive
21 nt. Unproductive
28 nt. Unproductive
37 nt. Unproductive
32 nt. Unproductive

KATMTVDKSSSTAYMELARLTSEDSAVYYCAR KATMTVDKSSSTAYMELARLTSEDSAVYYCAR 40 nt. 22 nt. Unproductive Unproductive

KATLTVDKSSSTAYMQLSSLTSEDSAVYYCVR KATLTVDTSSSTAYMQLSSLTSEDSAVYYCAR KATLTVDKPSSTAYMQLSSLTSEDSAVYYCAR KATLTVDKSSSTAYMQLSSLTSEDSAVYYCTL KATLTVDTSSSTAYMQLSSLTSEDSAVYYCAR KATLTVDKSSSTAYMQLSSLTSEDSAVYYCTI KATLTVDKSSSTAYMQLSSLTSEDSAVYYCAR KATLTVDTSSSTAYMQLSSLTSEDSAVYYCAR KATLTVDKSSSTAYMQLSSLTSEDSAVYYCAI KATLTVEKSSSTVYLELSRLTSDDSAVYYCAR KATLTVDKSSSTAYMQLSSLTSEDSAVYYCAK KATLTVDKSSSTAYMQLSSLTSEDSAVYYCAR KATLTVDKSSSTAYMQLSSLTSEDSAVYYCAP KAAWAVDTSSSTAYMQLSSLTSEDTAVYFCL* KATLTVDKPSDTAYMQLSSLTSEDSASYYCAR KATLTANTSSSTAYMELSSLTSEDSAVYYCAR KATLTVDTSSSTSYMQLSSLTSEDSAVYYCAR

RGLTYAMDY
YYSNYFDY
PNWDHYYYGMDV
LYYYAMDY
SSGYDY
GAARATNAY
GGFAY
SPMDY
EVPGGFYATDY
MDYYGSSLWFAY
TTVVAFDY
KRDYSTYFDH
TGTEFAY

TGTEFAY Ps.gene
24 nt. Ps.gene/Unproductiv
9 nt. Unproductive
23 nt. Unproductive
15 nt. Unproductive

RFTISRDNSQSILYLQMNALRAEDSATYYCAR RFTISRDNSQSILYLQMNALRAEDSATYYCAR RFTISRDNSQSILYLQMNALRAEDSATYYCAR

YMILGAMDY GYYYDGSYYAMDY 23 nt.

Unproductive

RFTISRDNAKNTLFLQMTSLRSEDTAMYYCAR RFIISRDNTKKTLYLQMSSLRSEDTALYYCAR RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAK RFIISRDNSKNTLYLQMNSLRAEDTAVYYCAK RFIISRDNSKKTLYLQMSSLRSEDTALYYCAR RFTISRDNSKNTLYLQMSSLRSEDTALYYCAK RFIISRDNTKKTLYLQMSSLRSEDTALYYCAR RFTISRDNAKNTLYLQMSSLRSEDTALYYCAR RFTISRDNAKNTLYLQMSSLKSEDTAMYYCAR

AKFHLYFDY

REGVVESRLDGDV

RGLHWFDP

RNYGSSPFDY

PPMMPSY

Ps.gene

PpMMPSY

Ps.gene

Ps.gene

43 nt. Ps.gene/Unproductiv
28 nt. Ps.gene/Unproductiv
35 nt. Unproductive

RFTISRDDSKSRVYLQMNSLRAEDTGIYYCTG

30 nt.

Unproductive

KATLTADKSSSTVYMELSRLTSEDSAVYFCAR

HEDRDSSGYAMDY



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CDR 2

FRAMEWORK 3

CDR 3

KABAT HUMAN VH1

STSTAYMELRSLRSEDTAVVYCAR GEGWDHFDY

HAQKFQG RVTIRRHKSTSTAYMELSSLRSEDTAVYYCAR GSRYGYDCSGYYYL

GYAQKFQG RVTMTRNTSISTATMELSSLRSEDTAVYYCAR LAHFSGSPVDWFDP

KABAT HUMAN VH2

KHQLQPSLKS RVTISVDTSKNQFSLKLSSVTAADTAVYYCAR GGVVPAAIMDV

KS RVTISVDTSKNQFSLKLSSVTAADTAVYYCAR MARYYDFWSGYSAYYDY

SLKS RLSISQDTSRNQFSLRLSSVTAADTAVYYCAR HRNWGSPVHFDY

ESTSTAYMELSSLRSEDTAVYYCAR DSYGDYGGHY

KABAT HUMAN VH3

ISYITSSSYTNYADSVKG RFTISRDNAKNSLYLQMNSLRADDTAVYYCAR DGRFGTYSPSDY
SVKG RFTISRDDSKSIAYLQVNSLKTEDTAVYYCTR TIYYDSSGYPYW

YADSVKG RFTISRDNAKNSLFLQMSSLRAEDTAFYYCAR GIALDAFDI

YYADSVRD RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAK 53 NT. UNPROD REARR DSVKG RFTISRDNAKNSLYLQMNSLRDEDTAVYYCAR DHSGTGGGGSGSYF

DSVKG RFTISRDNAKNSLYLQMNSLRDEDTAVYYCAR DHSGTGGGGSGSYF
VSAISGSGGSTYYADSVKG RFTISRDNPKNTLYLQMNSLRSEDTAVYYCAR KDNLWFDP

AVISYDGSNKYYADSVKG RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR DLGGRGVVVVPAPGGRSIYYYGMDV

GAVISYDGSNKYYADSVKG RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAS LEGIGTIYYYGMDV AKNSLYLQMNSLRAEDTAVYYCVR DDSSSWPKHFQH

QYAASVKG RFTISRDDSKNSLYLQMNSLNTEDTAVYYCVR SGVVPYLDY

KNOWN FAMILY

AVYYCAR DPRIAARPDYYYYMDV TAMYYCAR GAEVVEPTARYYYGLNV



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		•
FR1	CDR1_	FR2
YTFT	SYGIS	WVTTGPWTRDLRWMG
GEKPGSSVKVSCKASGYTFT	DYFMN	WMRQAPGQRLEWMG
QVQLQEIGPRTGEASETLSLICAVSGDSIS	SGNW*I	WVRQPPGKGLEWIG
QVQLQESGPGLVK*SETLSLTCTVSGGSIS	SYYWS	WIrqppGKGLEWIG
GYTFT	NYCMH	WVRQDHAQGLEWMG
QVQLQESGPGLVKpSETLSLYCAVSGDSIS	SGNW*I	WVRQPPGKGLEWIG
GPRLGEASETLSLTCTVSGGSIS	SSSYYw	WIRQPPGKGLEWIG
QVQLQESGPGLVKpSETLSLTCTVSGGSIS	SYYWS	WIRQPPGKGLEWIG
LSLICAVSGSSIS	SGNW*I	WVRQPPGKGLEWIG
SETLSLTCAVYGGSFS	GYYWS	WIRQPPGKGLEWIG
QVQLVQSGAEVKKPGASVKVSCKASGYTFT	NYCMH	WVRQVLAQGLEWMG
SETLSLICAVSGDSIS	SGNW*I	WVRQPPGKGLEWIG
SRAQTGEASETLSLTCTVSGGSIS	SSSYYWG	WIRQPPGKGLEWIG
CPLTCTVSGGSVSSGS	YYWS	WIRQPPGKGLEWIG
GLVKPSETLSLTCTVSGGSIS	SYYWS	WIGSPpGKGLEWIG
SFETLSLICAVSGDSIS	SGNW*I	WVRQPPGKGLEWIG
QVQLVQSGAEVKKPGSSVKVSCKASGGTFS	SYAIS	WVRQAPGQGLEWMG
QVQLQQWGAGLLKPSETLSLTCAVYGGSFS	GYYWS	WIRQPPGKGLEWIG
QLQLQESGPGLVKPSETLSLTCTVSGGSIS	SSSYYWG	WIRQPPGKGLEWIG
GPGLVKPSQTLSLTCTVSGGSIS	SGGYYWS	WIRQNPGKGLEWIG

- * indicates stop codon (unsure as sequence remains in frame)
 sequence termonates due to internal restriction site lower case denotes frame shift

CDR2	FR3	CDR3
[!] WISAYNGNTNYAQKLQG	RVTMTTDTSTSTAYMELRSLRSDDTAVYYCAR	DTVSS
WINAGNGNTKYSQKLQG	RVTITRDTSASTAYMQLSSLRSEDTAVYYCAR	DTVSS
EIHHSGSTYYNPSLKS	RITMSVDTSKNQFYLKLSS•	
RIYTSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSSVTAADTAVYYCAR	DTVSS
LVCPSDGSTSYAQKFQA	RVTITRDTSMSTAYMELSSLRSEDTAMYYCAR	DTVSS
EIHHSGSTYYNPSLKS	RITMSVDTSKNQFYLKLSS•	
EINHSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSS•	
YIYYSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSS•	
EIHHSGSTYYNPSLKS	RITMSVDTSKNQFYLKLSS•	
EINHSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSSVTAADTAVYYCAR	DTVSS
LVCPSDGSTSYAQKFQA	RVTITRDTSMSTAYMELSSLRSEDTAMYYCAR	DTVSS
EIHHSGSTYYNPSLKS	RITMSVDTSKNQFYLKLSS•	
SIYYSGSTYYNPSLKS	RVTIPVDTSKNQFSLKLSS•	
YIYYSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSSVTAADTAVYYCAR	DTVSS
RIYTSGSTNYNPSLKS	RVTMSVDTSKNQFSLKLSS•	
EIHHSGSTYYNPSLKS	RITMSVDTSKNQFYLKLSS•	
RIIPILGIANYAQKFQG	RVTITADKSTSTAYMELSSLRSEDTAVYYCAR	DTVS
EINHSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSS•	
EINHSGSTNYNPSLKS	RVTISVDTSKNQFSLKLSS•	
YIYYSGSTYYNPSLKS	RVTISVDTSKNQFSLKLSSVTAADTAVYYCAR	DTVSS



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pSW1

HindIII site AAGCTT

птп	arr	1 5	Tre	· AA	IGC I	1													
GCA	TGC	AAA	TTC	TAT			GAG	GAC!				AAA	TAC			CCI			
		1	0			20			30)		4	0			50			60
A GCT	_								P ACCA										
			0			80			90				0			.10			120
G GGA									S										
	-	13				40			150			16			1		- 0		180
S TCA									V GTT										
		19							210										240
L CTG									T 'ACA										
010		25		100		60	002		270			28		.001		90			300
S AGC	I ATC:	_	K						Q AAD									H	
11001	110	31		0,10		20			330		110		0	AI O		50	010		360
D GAT									R										
0711	J2 102	37		.100		80	101		390		non	40		AGG		10	IAC		420
Q CAA(-								י א אידי		ር አ ር ‹	ጉጥ አ	Sm		ccm	አ አ <i>ር</i>	CTIC	ሮ አ ኦ	ጥጥር
CAA																			



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pSW2

Hin	dII)	. AA	IGC 1	I.															
											М	ĸ	Y	L	L	P	т	Α	Α
GCA'	rgc <i>i</i>	AA.	rTCI	TAT	TC.	AAG	GAG	ACA	GTC	ATA		\AA'	CAC		ГТG	CCT			GCC
		10	_			20			30			4()			50			60
A	G	L	L	L	L	Α	Α	Q	P	Α	М	Α	Q	V	Q	L	Q	E	s
GCT(GGA:	rTG:	rta:	CTAC	CTC	GCT	GCC	CAA	CCA	GCG	ATG	SCC	CAG	GTG(CAG	CTG	CAG	GAG	TCA
		70)			80			90			100)		Τ	10			120
						,													
G	P	G·	L	V	Α	P	S	Q	S	L	S	I	T	С	T	V	S		F
GGA	CCT			GTG		CCC:	rca(AGC(CTG'	rcc	ATC/		rgcz		GTC 70	ľÇA	تاتاتا	11C
		130	J		1	40			130			100	,		-	, 0			
														_		_	_	_	
S	L	T	G	Y	G	V CEA	N	W	V GTT	R	Q TACC	P	P	G CA:	K NAC	G GCTI	L CTG		W TGC
I'CA'	TTA	190			1ی 2		AAC.		210		CAGC	220))	JGM		30	CIG	un u	240
_	_		_	7.7	_	5	<u></u>	NT	т	D	v	Νī	c	7\	т	ĸ	c	B	т.
	G GGA	M ATG	ላጥጥና ተ	w rgg(G GT	GAT(G GGA	AAC	ACA	GAC'	TAT?	AT:	rca(GCT					
010	00.2	25				60			270			28				90			300
s	I	s	ĸ	D	N	s	ĸ	s	Q	V	F	L	K	М	N	s	L	Н	т
		AGC	AAG	GAC	AAC	TCC	AAG	AGC	CAA	GTT'	TTC:	rta.	AAA	ATG	AAC	AGT	CTG	CAC	ACT
		31	0		3	20			330			34)		3	50			360
D	D	T	Α	R	Y	Y	С	Α	R	E	R		Y			D			
GAT	GAC			AGG:			rgr(GCC	AGA(GAG.	AGA	3AT' 40		AGG		'GAC' 10	TAC	TGG	GGC 420
		37	J		3	80			390			40	J		. 4	10			420
Q	G	T	T	V	Т	V	S	S			C 7 C /	2m.C.	~ 7 7 1	mmc	~~~	' A A C	~mm	~~ n	mcc
CAA	GGC	4CC 43		GTC		GTC:	I.C.C.	TCA	TAA:	ľAA	GAG	46		TTC		.AAG .70	CII	GCA	480
		40	•		-	40			150				•		_	•			
												-	_	_	_	-	70	78	~
<i>ת</i> ת ת	יים יים	ייזי איז	יייר	A A C (220	מרמנ	<u>ст</u> С:	ביד ב	M .ATG							A GCA			G 'GGA
AAA	110.	49		AAG		00	310	AI.	510		Inco	52		001		30	-	-	540
τ.	т	т	т	7	7\	0	מ	7	М	7\	ח	т	17	Τ.	т	0	ς	P	Δ
ኍ ፐፐG	TTA'	ь ГТА	L CTC	A GCT	GCC	CAA	CCA	GCG	ATG	GCC	GAC	ATC	V GTC	CTG	ACT	'CĀG	TCT	CCA	.GCC
		55				60			570			58				90			600
S	т.	S	Δ	S	V	G	F.	т	V	т	I	т	С	R	Α	s	G	N	I
TCC	CTT'	rct.	GCG'	rcr(GTG	GGA	GAA	ACI	GTC	ACC.	ATC	ACA'	TGT	CGA	GCA	AGT	GGG	AAT	'ATT
		61	0		6	20			630			64	0		6	50			660
Н	N	Y	L	Α	W	Y	Q	Q	K	Q	G	ĸ	s	P	Q	L	L	V	Y
	AAT'	TAT'	TTA	GCA'	rgg	TAT	CAG	CAG	AAA	CAG	GGA	AAA	TCT	CCT	CAG	CTC	CTG	GTC	TAT
		67	0		6	80		,	690			70	O		7	10			720

720



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Y T T T L A D G V P S R F S G S G S G T TATACAACAACCTTAGCAGATGGTGTGCCATCAAGGTTCAGTGGCAGTGGATCAGGAACA 730 740 750 760 770 780

Q Y S L K I N S L Q P E D F G S Y Y C Q CAATATTCTCTCAAGATCAACAGCCTGCAACCTGAAGATTTTGGGAGTTATTACTGTCAA 790 800 810 820 830 840

H F W S T P R T F G G G T K L E I K R CATTTTTGGAGTACTCCTCGGACGTTCGGTGGAGGCACCAAGCTGGAAATCAAACGGTAA 850 860 870 880 890 900

TAAGAGCTCGAATTC 910

FIG. 14 b

pSW1HPOLYMYC

HindIII site AAGCTT

M K Y L L P T A A GCATGCAAATTCTATTTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACGGCAGCC 10 20 30 40 50 60

A G L L L L A A Q P A M A Q V Q L Q GCTGGATTGTTATTACTCGCTGCCCAACCAGCGATGGCCCAGGTGCAG**CTGCAG** 70 80 90 100 110 PstI

Polylinker TCTAGA GTCGAC CTCGAG XbaI SalI XhoI

MYC PEPTIDE

V T V S S E O K L I S E E D L N * *

GGTCACCGTCTCCTCAGAACAAAACTCATCTCAGAAGAGGATCTGAATTAATAA

BstEII

GGGCTAAGCTCGAATTC



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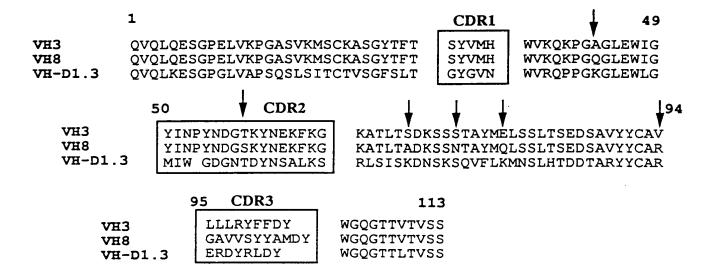


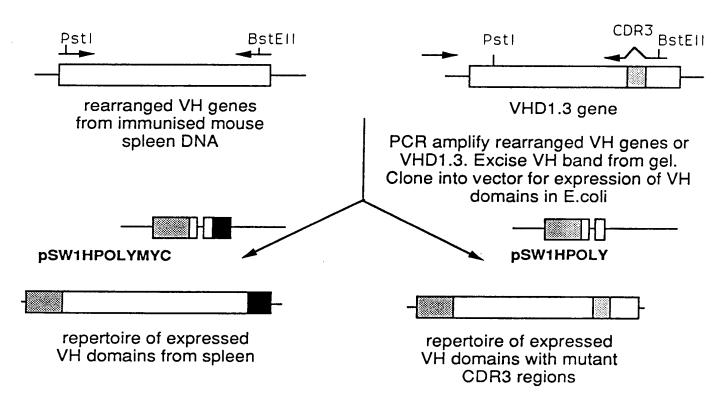
FIG. 16

FR1	QVQLQESGGGLVQPGGSLRLSCAASGFTFS	
	SYAMS	CDR1
FR2	WVRQAPGKGLEWVS	
	AISGSGGSTYYADSVKG	CDR2
FR3	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAM	
	WRGIATPVSFDLGYFDY	CDR3

FIG. 17



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Assay for binding to antigen



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et al

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pSW2HPOLY

HindIII AAGCTT

MKYLLPTAA GCATGCAAATTCTATTTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACGGCAGCC 50 20 30 40

AGLLLLAAQPAMAQVQLQ GCTGGATTGTTATTACTCGCTGCCCAACCAGCGATGGCCCAGGTGCAGCTGCAG 80 90 100 110 PstI 70

TCTAGA GTCGAC CTCGAG

XbaI SalI

XhoI

V T V S S **GGTCACC**GTCTCCTCATAATAAGAGCTCGAATTCGCCAAGCTTGCATGC BstEII 430 440 450 460 470

M K Y L L P T A A A G AAATTCTATTTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACGGCAGCCGCTGGA 500 510 520 530

480

L L L L A A Q P A M A D I V L T Q S P A TTGTTATTACTCGCTGCCCAACCAGCGATGGCCGACATCGTCCTGACTCAGTCTCCAGCC 560 570 580 590 550

S L S A S V G E T V T I T C R A S G N I TCCCTTTCTGCGTCTGTGGGAGAAACTGTCACCATCACATGTCGAGCAAGTGGGAATATT 620 630 640

H N Y L A W Y Q Q K Q G K S P Q L L V Y CACAATTATTTAGCATGGTATCAGCAGAAACAGGGAAAATCTCCTCAGCTCCTGGTCTAT 670 680 690 700 710

Y T T T L A D G V P S R F S G S G S G T TATACAACAACCTTAGCAGATGGTGTGCCATCAAGGTTCAGTGGCAGTGGATCAGGAACA 730 740 750 760 770

Q Y S L K I N S L Q P E D F G S Y Y C Q CAATATTCTCTCAAGATCAACAGCCTGCAACCTGAAGATTTTGGGAGTTATTACTGTCAA 790 800 810 820 830

H F W S T P R T F G G G T K L E I K R CATTTTTGGAGTACTCCTCGGACGTTCGGTGGAGGCACCAAGCTGGAAATCAAACGGTAA 850 860 870 880

TAAGAGCTCGAATTC 910



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MKYLLPT AAGCTTGCATGCAAATTCTATTTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACG 20 30 40 50 A A A G L L L A A Q P A M A Q V Q L Q GCAGCCGCTGGATTGTTATTACTCGCTGCCCAACCAGCGATGGCCCAGGTGCAGCTGCAG 100 E S G P G L V A P S O S L S I T C T V S GAGTCAGGACCTGGCCTGGTGGCGCCCTCACAGAGCCTGTCCATCACATGCACCGTCTCA 140 150 160 170 G F S L T G Y G V N W V R Q P P G K G L GGGTTCTCATTAACCGGCTATGGTGTAAACTGGGTTCGCCAGCCTCCAGGAAAGGGTCTG 200 210 220 E W L G M I W G D G N T D Y N S A L K S GAGTGGCTGGGAATGATTTGGGGTGATGGAAACACAGACTATAATTCAGCTCTCAAATCC 260 270 280 R L S I S K D N S K S Q V F L K M N S L AGACTGAGCATCAGCAAGGACAACTCCAAGAGCCAAGTTTTCTTAAAAATGAACAGTCTG 330 H T D D T A R Y Y C A R E R D Y R L D Y 380 390 400 $\hbox{\tt W} \quad \hbox{\tt G} \quad \hbox{\tt O} \quad \hbox{\tt G} \quad \hbox{\tt T} \quad \hbox{\tt T} \quad \hbox{\tt V} \quad \hbox{\tt T} \quad \hbox{\tt V} \quad \hbox{\tt S} \quad \hbox{\tt S} \quad \hbox{\tt G} \quad \hbox{\tt G} \quad \hbox{\tt G} \quad \hbox{\tt A} \quad \hbox{\tt P} \quad \hbox{\tt A} \quad \hbox{\tt A} \quad \hbox{\tt P}$ TGGGGCCAAGGCACCACGGTCACCGTCTCCTCAGGTGGTGCTCCAGCAGCTGCACCT 440 450 460 470 A G G G Q V Q L K E S G P G L V A P S O GCTGGAGGAGGACAGGTGCAGCTGAAGGAGTCAGGACCTGGCCTGGTGGCGCCCTCACAG 500 510 520 530 S L S I T C T V S G F S L T G Y G V N W AGCCTGTCCATCACATGCACCGTCTCAGGGTTCTCATTAACCGGCTATGGTGTAAACTGG 560 570 580 V R Q P P G K G L E W L G M I W G D G N GTTCGCCAGCCTCCAGGAAAGGGTCTGGAGTGGCTGGGAATGATTTGGGGTGATGGAAAC 620 630 640 650 TDYNSALKSRLSISKDNSKS ACAGACTATAATTCAGCTCTCAAATCCAGACTGAGCATCAGCAAGGACAACTCCAAGAGC 690 700 Q V F L K M N S L H T D D T A R Y Y C A CAAGTTTTCTTAAAAATGAACAGTCTGCACACTGATGACACAGCCAGGTACTACTGTGCC 740 750 RERDYRLDYWGQGTTVTV AGAGAGAGAGATTATAGGCTTGACTACTGGGGCCAAGGCACCACGGTCACCGTCTCCTCA 800 810 820 830 TAATAAGAGCTC 850



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M K Y L L P T A A GCATGCAAATTCTATTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACGGCAGCC 10 20 30 40 50 60

A G L L L L A A Q P A M A Q V Q L Q E S GCTGGATTGTTATTACTCGCTGCCCAACCAGCGATGGCCCAGGTGCAGCTGCAGGAGTCA 70 80 90 100 110 120

G P G L V A P S Q S L S I T C T V S G F GGACCTGGCCTGGTGGCCCCTCACAGAGCCTGTCCATCACATGCACCGTCTCAGGGTTC 130 140 150 160 170 180

S L T G Y G V N W V R Q P P G K G L E W TCATTAACCGGCTATGGTGTAAACTGGGTTCGCCAGCCTCCAGGAAAGGGTCTGGAGTGG 190 200 210 220 230 240

L G M I W G D G N T D Y N S A L K S R L CTGGGAATGATTTGGGGTGATGGAAACACAGACTATAATTCAGCTCTCAAATCCAGACTG 250 260 270 280 290 300

S I S K D N S K S Q V F L K M N S L H T AGCATCAGCAAGGACAACTCCAAGAGCCAAGTTTTCTTAAAAATGAACAGTCTGCACACT 310 320 330 340 350 360

D D T A R Y Y C A R E R D Y R L D Y W G GATGACAGCCAGGTACTGCTGCCAGAGAGAGAGATTATAGGCTTGACTACTGGGGC 370 380 390 400 410 420

A A Q G D I T A P G G A R R L T G D Q T GCTGCTCAGGGCGATATTACTGCACCCGGCGGTGCTCGCCGTTTAACGGGTGATCAGACT 490 500 510 520 530 540

A A L R D S L S D K P A K N I I L L I G GCCGCTCTGCGTGATTCTCTTAGCGATAAACCTGCAAAAAATATTATTTTGCTGATTGGC 550 560 570 580 590 600

D G M G D S E I T A A R N Y A E G A G G GATGGGATGGGGGAAATTACTGCCGCACGTAATTATGCCGAAGGTGCGGCGGC 610 620 630 640 650 660

K K T G K P D Y V T D S A A S A T A W S AAAAAAACCGGCAAACCGGACTACGTCACCGACTCGGCTGCATCAGCAACCGCCTGGTCA 730 740 750 760 770 780

FIG. 21 a

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T G V K T Y N G A L G V D I H E K D H P ACCGGTGTCAAAACCTATAACGGCGCGCTGGGCGTCGATATTCACGAAAAAGATCACCCA 790 800 810 820 830 840

T I L E M A K A A G L A T G N V S T A E ACGATTCTGGAAATGGCAAAAGCCGCAGGTCTGGCGACCGGTAACGTTTCTACCGCAGAG 850 860 870 880 890 900

L Q D A T P A A L V A H V T S R K C Y G TTGCAGGATGCCACGCCCGCTGCGCTGGTGGCACATGTGACCTCGCGCAAATGCTACGGT 910 920 930 940 950 960

P S A T S E K C P G N A L E K G G K G S CCGAGCGCGACCAGTGAAAAATGTCCGGGTAACGCTCTGGAAAAAGGCGGAAAAGGATCG 970 980 990 1000 1010 1020

F A E T A T A G E W Q G K T L R E Q A Q TTTTGCTGAAACGCCACCGCTGGTGAATGGCAGGGAAAAACGCTGCGTGAACAGGCACAG 1090 1100 1110 1120 1130 1140

A R G Y Q L V S D A A S L N S V T E A N GCGCGTGGTTATCAGTTGGTGAGCGATGCTGCCTCACTGAATTCGGTGACGGAAGCGAAT 1150 1160 1170 1180 1190 1200

Q Q K P L L G L F A D G N M P V R W L G CAGCAAAAACCCCTGCTTGGCCTGTTTGCTGACGGCAATATGCCAGTGCGCTGGCTAGGA 1210 1220 1230 1240 1250 1260

P K A T Y H G N I D K P A V T C T P N P CCGAAAGCAACGTACCATGGCAATATCGATAAGCCCGCAGTCACCTGTACGCCAAATCCG 1270 1280 1290 1300 1310 1320

Q R N D S V P T L A Q M T D K A I E L L CAACGTAATGACAGTGTACCAACCCTGGCGCAGATGACCGACAAAGCCATTGAATTGTTG 1330 1340 1350 1360 1370 1380

S K N E K G F F L Q V E G A S I D K Q D AGTAAAAATGAGAAAGGCTTTTTCCTGCAAGTTGAAGGTGCGTCAATCGATAAACAGGAT 1390 1400 1410 1420 1430 1440

H A A N P C G Q I G E T V D L D E A V Q CATGCTGCGAATCCTTGTGGGCAAATTGGCGAGACGGTCGATCTCGATGAAGCCGTACAA 1450 1460 1470 1480 1490 1500

R A L E F A K K E G N T L V I V T A D H CGGGCGCTGGAATTCGCTAAAAAGGAGGGTAACACGCTGGTCATAGTCACCGCTGATCAC 1510 1520 1530 1540 1550 1560



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A H A S Q I V A P D T K A P G L T Q A L GCCCACGCCAGCCAGATTGTTGCGCCGGATACCAAAGCTCCGGGCCTCACCCAGGCGCTA 1570 1580 1590 1600 1610 1620

N T K D G A V M V M S Y G N S E E D S Q AATACCAAAGATGGCGCAGTGATGGTGATGAGTTACGGGAACTCCGAAGAGGATTCACAA 1630 1640 1650 1660 1670 1680

E H T G S Q L R I A A Y G P H A A N V V GAACATACCGGCAGTCAGTTGCGTATTGCGGCGTATGGCCCGCATGCCGCCAATGTTGTT 1690 1700 1710 1720 1730 1740

G L T D Q T D L F Y T M K A A L G L K * GGACTGACCGACCGATCTCTTCTACACCATGAAAGCCGCTCTGGGGCTGAAATAA 1750 1760 1770 1780 1790 1800

AACCGCGCCCGGGAGTGAATTTTCGCTGCCGGGTGGTTTTTTTGCTGTTAGC 1810 1820 1830 1840 1850

FIG. 21c



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MKYLLPTAA GCATGCAAATTCTATTTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACGGCAGCC 10 20 30 40 50

A G L L L A A Q P A M A Q V Q L Q E S GCTGGATTGTTATTACTCGCTGCCCAACCAGCGATGGCCCAGGTGCAGCTGCAGGAGTCA 100 110 70 80 90

G P G L V A P S Q S L S I T C T V S G F GGACCTGGCCTGGCGCCCCTCACAGAGCCTGTCCATCACATGCACCGTCTCAGGGTTC 150 160 170 180 130 140

S L T G Y G V N W V R Q P P G K G L E W TCATTAACCGGCTATGGTGTAAACTGGGTTCGCCAGCCTCCAGGAAAGGGTCTGGAGTGG 190 200 210 220 230

L G M I W G D G N T D Y N S A L K S R L CTGGGAATGATTTGGGGTGATGGAAACACAGACTATAATTCAGCTCTCAAATCCAGACTG 290 250 260 270 280

S I S K D N S K S Q V F L K M N S L H T AGCATCAGCAAGGACAACTCCAAGAGCCAAGTTTTCTTAAAAATGAACAGTCTGCACACT 310 320 330 340 350

D D T A R Y Y C A R E R D Y R L D Y W G GATGACACAGCCAGGTACTACTGTGCCAGAGAGAGAGATTATAGGCTTGACTACTGGGGC 410 370 380 390 400

OGTTVTVSS** 460 470 480 430 440 450

M K Y L L P T A A A G L TTCTATTTCAAGGAGACAGTCATAATGAAATACCTATTGCCTACGGCAGCCGCTGGATTG 490 500 510 520 530

LLLAA'OPAMADIELVDLEIK TTATTACTCGCTGCCCAACCAGCGATGGCCGACATCGAGCTCGTCGACCTCGAGATCAAA 580 590 550 560 570

REQKLISEEDLN** CGGGAACAAAACTCATCTCAGAAGAGGATCTGAATTAATAATGATCAAACGGTAATAAG 610 620 630 640 650

GATCCAGCTCGAATTC 670



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Q V Q L Q E S G P G L V Q P S Q S L S I CAGGTGCAGCTGCAGGAGTCAGGACCTGGCCTAGTGCAGCCCTCACAGAGCCTGTCCATC G N T C T V S G F S L T S Y G V H W V R Q S ACCTGCACAGTCTCTGGTTTCTCATTAACTAGCTATGGTGTACACTGGGTTCGCCAGTCT С P G K G L E W L G M I W G D G N T D Y N CCAGGAAAGGGTCTGGAGTGGCTGGGAATGATTTGGGGTGATGGAAACACAGACTATAAT S A L K S R L S I S K D N S K S Q V F L TCAGCTCTCAAATCCAGACTGAGCATCAGCAAGGACAACTCCAAGAGCCAAGTTTTCTTA K M N S L H T D D T A R Y Y C A R E R D AAAATGAACAGTCTGCACACTGATGACACAĞCCAGGTACTACTGTGCCAGAGAGAGAGAT

FIG. 23

Y R L D Y W G Q G T T V T V S S TATAGGCTTGACTACTGGGGCCAAGGGACCACGGTCACCGTCTCCTCA